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22850 7590 08/05/2009 OBLON, SPIVAK, MCCLELLAND MAIER & NEUSTADT, P.C. 1940 DUKE STREET			EXAMINER	
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			1657	
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# Please find below and/or attached an Office communication concerning this application or proceeding.

The time period for reply, if any, is set in the attached communication.

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patentdocket@oblon.com oblonpat@oblon.com jgardner@oblon.com

	Application No.	Applicant(s)
	09/990,049	FORD ET AL.
Office Action Summary	Examiner	Art Unit
	David M. Naff	1657
The MAILING DATE of this communication app Period for Reply	ears on the cover sheet with the c	orrespondence address
A SHORTENED STATUTORY PERIOD FOR REPLY WHICHEVER IS LONGER, FROM THE MAILING DA  - Extensions of time may be available under the provisions of 37 CFR 1.13 after SIX (6) MONTHS from the mailing date of this communication.  - If NO period for reply is specified above, the maximum statutory period w.  - Failure to reply within the set or extended period for reply will, by statute, Any reply received by the Office later than three months after the mailing earned patent term adjustment. See 37 CFR 1.704(b).	ATE OF THIS COMMUNICATION 36(a). In no event, however, may a reply be tim vill apply and will expire SIX (6) MONTHS from cause the application to become ABANDONE	lely filed the mailing date of this communication. (35 U.S.C. § 133).
Status		
Responsive to communication(s) filed on 14 Ma     This action is <b>FINAL</b> . 2b)☑ This     Since this application is in condition for allowar closed in accordance with the practice under E	action is non-final. nce except for formal matters, pro	
Disposition of Claims		
4) ☐ Claim(s) 25-33,35-43,45,47 and 67 is/are pend 4a) Of the above claim(s) 40-42,47 and 67 is/are 5) ☐ Claim(s) is/are allowed. 6) ☐ Claim(s) 25-33, 35-39, 43 and 45 is/are rejecte 7) ☐ Claim(s) is/are objected to. 8) ☐ Claim(s) are subject to restriction and/or	re withdrawn from consideration.	
Application Papers		
9) The specification is objected to by the Examine  10) The drawing(s) filed on is/are: a) access Applicant may not request that any objection to the of Replacement drawing sheet(s) including the correction in the original than the correction of the correction of the original than the correction of the correcti	epted or b) objected to by the Eddrawing(s) be held in abeyance. See ion is required if the drawing(s) is obj	e 37 CFR 1.85(a). ected to. See 37 CFR 1.121(d).
Priority under 35 U.S.C. § 119		
12) Acknowledgment is made of a claim for foreign a) All b) Some * c) None of:  1. Certified copies of the priority documents 2. Certified copies of the priority documents 3. Copies of the certified copies of the prior application from the International Bureau * See the attached detailed Office action for a list of	s have been received. s have been received in Application ity documents have been receive I (PCT Rule 17.2(a)).	on No ed in this National Stage
Attachment(s)  1) \[ \sum \text{Notice of References Cited (PTO-892)} \]	4) 🗖 Intonious Summans	(PTO 413)
<ol> <li>Notice of References Cited (PTO-892)</li> <li>Notice of Draftsperson's Patent Drawing Review (PTO-948)</li> <li>Information Disclosure Statement(s) (PTO/SB/08)         Paper No(s)/Mail Date     </li> </ol>	4)  Interview Summary Paper No(s)/Mail Da 5)  Notice of Informal P 6)  Other:	ite

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#### **DETAILED ACTION**

Claims in the application are 25-33, 35-43, 45, 47 and 67.

A response of 5/14/09 to a restriction requirement of 4/17/09 elected Group I claims 25-33, 35-39, 43 and 45 with traverse.

The traverse is on the ground that requirements for distinctness specified in MPEP 806.05(j) have not been met. The traverse points out that claim 25 is directed to producing a metal particle nucleic acid composite while claim 40 is directed to producing a nanowire which comprises providing a metal particle nucleic acid composite. However, in claim 25 bases of a nucleic acid are reacted with a specific metal complex to form a metal complex-nucleic acid conjugate, whereas claim 40 requires attaching an imidazole group as a metal ligand to a cytosine of a polynucleotide, and metalating the attached imidazole with a metal complex having a tridentate ligand and a leaving group to form a conjugated metal complex. These are mutually exclusive methods that are not obvious variants and which have different modes of operation. Therefore, the requirements for distinctness are met. The traverse further points out that claim 67 depends on previously examined claim 40. However, claim 40 previously examined was claim 40 prior to the amendment of 1/29/09. Amended claim 40 in the amendment of 1/29/09 has not been previously examined. The restriction requirement is still considered proper, and is adhered to and made final.

Claims 40-42, 47 and 67 are withdrawn from further consideration pursuant to 37 CFR 1.142(b), as being drawn to a nonelected invention, there being no allowable generic or linking claim. Applicant timely traversed the restriction (election) requirement in the reply filed on 5/14/09.

Claims examined on the merits are 25-33, 35-39, 43 and 45.

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### Claim Rejections - 35 USC § 112

The following is a quotation of the first paragraph of 35 U.S.C. 112:

The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.

Claims 25-33, 35-39, 43 and 45 are rejected under 35 U.S.C. 112, first paragraph, as failing to comply with the written description requirement. The claim(s) contains subject matter which was not described in the specification in such a way as to reasonably convey to one skilled in the relevant art that the inventor(s), at the time the application was filed, had possession of the claimed invention.

Description is not found in the specification of producing the claimed composite containing metal particles of only sub-nanometer size. The specification (page 15, line 7) discloses a total thickness between 3 nm and 6 nm. Since DNA is 2 nm thick (specification page 4, line 19), the metal particles will be 1-4 nm in size. No description is seen in the specification of controlling the process to produce metal particles of only sub-nanometer size. The specification indicates the DNA contains metal particles varying in size, and Figures 2-6, 13 and 14 show total thickness of the metalized DNA varying along its length, which indicates that metal particle size varies along the length. The specification fails to support that all metal particles on the DNA are sub-nanometer in size.

## Response to Arguments

The amendment of 1/29/09 refers to the specification (page 4, lines 24-27) disclosing that in contrast to the procedure of Pompe et al, the sub-nanometer size of the platinum particles in the nanoparticle/DNA composite produced according to the present invention are stable in time, at least for weeks or months. The amendment also cites Figures 2 and 5 as further supporting a sub-nanometer size. However, other portions of the specification indicate the metal particles

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are not less than one nanometer in size. Furthermore, the statement at page 4, lines 24-27 is in regard to platinum particles, and does not to any metal particles as encompassed by the claims. The specification fails to disclose that Figures 2 and 5 show metal particles that are only less than 1 nanometer in size. The specification discloses (page 16, last two lines of the last paragraph (Example 3) and page 17, Example 5) that Figures 2 and 5 show images of elongated segments of DNA without any evidence of Pt-particles. It is uncertain how showing elongated segments of DNA without Pt-particles supports metal particles of only less than 1 nanometer size. The specification discloses that DNA is 2 nm thick (page 4, line 19), which indicates that metal particles will be 1-4 nm in size when total thickness is between 3 nm and 6 nm as disclosed in the specification. Description is not found in the specification of controlling the process to produce metal particles of only less than 1 nanometer size.

## Claim Rejections - 35 USC § 103

Claims 25-31, 33, 35-39, 43 and 45 are rejected under 35 U.S.C. 103(a) as being unpatentable over Pompe et al (AR) in view of Singh et al (5,560,960) and Richter et al (AQ) for reasons in the previous office action of 7/29/08, and for reasons herein.

The claims are drawn to a process of producing a metal particle nucleic acid composite containing metal particles by reacting a nucleic acid specific metal complex with a nucleic acid to produce a metal complex-nucleic acid conjugate, removing non-conjugated metal complexes and/or non-conjugated byproducts, and reacting the conjugate with a reducing agent to produce the metal particle nucleic acid composite. The metal complex-nucleic acid conjugate is formed by the specific reacting of the nucleic acid specific metal complex with bases of the nucleic acid, the metal particle-nucleic acid composite is catalytically active towards electroless metallization, the metal particles in the composite are not visualized by atomic force microscopy, and the

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metal particles in the composite are sub-nanometer in size. Also claimed is a metal particlenucleic acid composite resulting from the process.

Pompe et al disclose (page 1090, left col, second full paragraph) that Pt(II) and Pd(II) complexes such as cis-diamminedichloroplatinum attach to DNA bases to form stable monofunctional and bifunctional adducts. Further disclosed (third full paragraph of the left col) is that the Pt-DNA bond is not broken during reduction, and that Pt(II) and Pd(II) complexes attached to DNA double chain can act as nucleation centers for the growth of metal clusters. Also disclosed is carrying out metallization of DNA by adding DNA to Pd salt solution followed by adding a reducing agent, and obtaining clusters on the DNA of 3 to 5 nm in diameter in a few seconds after adding the reducing agent (paragraph bridging the cols, page 1090). Further disclosed is that a wide spread of cluster size distribution occurs reaching from less than 1 nm to more than 20 nm (page 1086, right col, line 8 from the bottom), and obtaining an average size of 1.9 nm (page 1087, left col, lines 1-4).

Singh et al disclose (paragraph bridging cols 1 and 2) precipitating nanometer-sized metal particles from solution within vesicles made from polymerizable phospholipids.

Polymerized phospholipids are formed and added to a electroless plating solution. Before the electroless plating solution is added, palladium or platinum is provided on the inside surface of vesicles to function as a catalyst (col 3, lines 44-64). To insure that metal particles form only on the inside surface, any metal on the exterior surface of the vesicle is removed such as by using a chelating agent and gel filtering, or by passing the vesicles through an ion exchange column.

Singh et al further disclose (col 5, line 18) using cobalt, nickel or iron when producing metal nanoparticles by electroless plating.

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Richter et al disclose (page 508 and 510) metallization of DNA similar to Pompe et al and disclose formation of clusters of 1-5 nm diameter on DNA (page 508, left col, third full paragraph).

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It would have been obvious to attach cis-diamminedichloroplatinum to DNA as disclosed by Pompe et al, and then use a reducing agent to obtain DNA containing attached platinum metal catalysis for use in electroless deposition of metal on the DNA as suggested by Singh et al subjecting vesicles containing Pd or Pt to electroless metal deposition and as suggested by Pompe et al carrying out metallization of DNA by treating a DNA solution with a Pd salt solution, and then adding a reducing agent to form metal clusters on the DNA. Removing any nonattached metal complex from the DNA before electroless metallization would have been obvious to prevent the non-attached metal complex from forming metal particles as suggested by Singh et al removing metal from the exterior of vesicles to prevent metal particles from being formed on the vesicles exterior surface. The objective of Pompe et al is to obtain metal clusters on the DNA and not at other places, and to accomplish this one would obviously have to remove nonattached metal complex before electroless metallization. Removing any non-conjugated byproducts would have been obvious simply to prevent any possible inference with subsequent reactions. It would have been apparent from Richter et al that metal clusters of 1-5 nm diameter can be obtained, and it would have been obvious to produce clusters not thicker than DNA since this is an objective of Pompe et al (page 1090, left col, first full paragraph). Such clusters will not be capable of being visualized by atomic force microscopy. Additionally, Pompe et al disclose cluster sizes varying from less than 1 nm to above sub-nanometer, and an average size of 1.9 nm. This would result in the some clusters of less than 1 nm in size being present. Reacting DNA with cis-diamminedichloroplatinum as disclosed by Pompe et al followed by reducing as set forth above will inherently result in metallization of bases, and provide a metal

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nanoparticle active towards electroless metallization. The metallization of Pompe et al and Richter et al is controlled since they disclose controlling the time of metallization to control the size of clusters. The use of cobalt, nickel or iron when producing metal nanoparticles by electroless plating as disclosed by Singh et al would have suggested using a electroless plating solution as in claim 38.

### Response to Arguments

The amendment of 1/29/09 urges that examination determining obviousness must comply with the rationale specified in *KSR International Co. v. Teleflex Inc.* However, the reasons for obviousness, based on the disclosures of the references, set forth above provides the rationale required. As noted above, the disclosure of sub-nanometer platinum particles as being in contrast to Pompe et al is inconsistent with another portion of the specification that indicate metal particles are 1-4 nm in size. Pompe et al disclose cluster sizes varying from less than 1 nm to above 20 nm, and an average size of 1.9 nm. This will result in metal particles of less than 1 nm in size. The present specification fails to establish that the invention produces a composite having only sub-nanometer metal particles. The working examples in the specification do not show producing a composite containing only metal particles less than 1 nm in size. Claim 40 is a withdrawn non-elected invention, and comments regarding the obviousness of claim 40 are moot.

## Claim Rejections - 35 USC § 103

Claim 32 is rejected under 35 U.S.C. 103(a) as being unpatentable over the references as applied to claims 25-31, 33, 35-39, 43 and 45 above, and further in view of Newsman et al (5,670,680) for reasons in the previous office action, and for reasons herein.

The claim requires a gaseous reducing agent.

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Singh et al disclose using hydrogenation (col 4, line 57) for reducing metal ions to produce metals in a process of producing metal nanoparticles by electroless plating.

Newman et al disclose using hydrogen gas for hydrogenation in producing metal complexes.

It would have been obvious to use hydrogen gas as a reducing agent to reduce the metal of a conjugate of a metal complex and DNA disclosed by Pompe et al as suggested by Singh et al and Newman et al.

### Response to Amendment

The amendment urges that claim 32 depends from claim 25, and Newman et al do not cure the deficiency of the references applied to claim 25. However, as set forth above, the references applied to claim 25 are not deficient.

### **Double Patenting**

The nonstatutory double patenting rejection is based on a judicially created doctrine grounded in public policy (a policy reflected in the statute) so as to prevent the unjustified or improper timewise extension of the "right to exclude" granted by a patent and to prevent possible harassment by multiple assignees. A nonstatutory obviousness-type double patenting rejection is appropriate where the conflicting claims are not identical, but at least one examined application claim is not patentably distinct from the reference claim(s) because the examined application claim is either anticipated by, or would have been obvious over, the reference claim(s). See, e.g., *In re Berg*, 140 F.3d 1428, 46 USPQ2d 1226 (Fed. Cir. 1998); *In re Goodman*, 11 F.3d 1046, 29 USPQ2d 2010 (Fed. Cir. 1993); *In re Longi*, 759 F.2d 887, 225 USPQ 645 (Fed. Cir. 1985); *In re Van Ornum*, 686 F.2d 937, 214 USPQ 761 (CCPA 1982); *In re Vogel*, 422 F.2d 438, 164 USPQ 619 (CCPA 1970); and *In re Thorington*, 418 F.2d 528, 163 USPQ 644 (CCPA 1969).

A timely filed terminal disclaimer in compliance with 37 CFR 1.321(c) or 1.321(d) may be used to overcome an actual or provisional rejection based on a nonstatutory double patenting ground provided the conflicting application or patent either is shown to be commonly owned with this application, or claims an invention made as a result of activities undertaken within the scope of a joint research agreement.

Effective January 1, 1994, a registered attorney or agent of record may sign a terminal disclaimer. A terminal disclaimer signed by the assignee must fully comply with 37 CFR 3.73(b).

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Claims 25-33, 35-39, 43 and 45 are rejected under the judicially created doctrine of obviousness-type double patenting as being unpatentable over claims 1-32 of U.S. Patent 6,884,587 in view of Singh et al.

The claims of the patent require metallization of a nucleic acid to produce a metal nanoparticle-nucleic acid composite.

It would have been obvious in view of Singh et al for the type of reasons set forth above to remove non-conjugated metal complexes and/or non-conjugated by-products, if formed, before treatment with a reducing agent in the process of the copending application claims for metallization of DNA. The presence of extraneous metal complex or other by-products will obviously be a contaminant that can interfere with subsequent reactions.

### Response to Arguments

The amendment urges this rejection be held in abeyance until identification of patentable subject matter in this application. However, this rejection cannot be withdrawn until overcome.

#### Conclusion

Any inquiry concerning this communication or earlier communications from the examiner should be directed to David M. Naff whose telephone number is 571-272-0920. The examiner can normally be reached on Monday-Friday 9:30-6:00.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Jon Weber can be reached on 571-272-0925. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

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Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see <a href="http://pair-direct.uspto.gov">http://pair-direct.uspto.gov</a>. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

/David M. Naff/ Primary Examiner, Art Unit 1657

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DMN 9/29/09